

GenCore version 5.1.6
Copyright (c) 1993 - 2003 Compugen Ltd.

OM protein - protein search, using sw model

Run on: August 9, 2003, 16:11:13 ; Search time 53.7429 Seconds
(without alignments)
56.115 Million cell updates/sec

Title: US-09-905-691-2

Perfect score: 19
Sequence: 1 ARAARRAARAAARRAEEA 19

Scoring table: OLIGO
Gapop 60.0 , Gapext 60.0

Searched: 1107863 seqs, 158746573 residues

Word size : 0
Total number of hits satisfying chosen parameters: 1107863

Minimum DB seq length: 0
Maximum DB seq length: 2000000000

Post-processing: Listing first 45 summaries

Database : A_Geneseq_19Jun03.*

```
1: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA1980.DAT.*
2: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA1981.DAT.*
3: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA1982.DAT.*
4: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA1983.DAT.*
5: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA1984.DAT.*
6: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA1985.DAT.*
7: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA1986.DAT.*
8: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA1987.DAT.*
9: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA1988.DAT.*
10: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA1989.DAT.*
11: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA1990.DAT.*
12: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA1991.DAT.*
13: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA1992.DAT.*
14: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA1993.DAT.*
15: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA1994.DAT.*
16: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA1995.DAT.*
17: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA1996.DAT.*
18: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA1997.DAT.*
19: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA1998.DAT.*
20: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA1999.DAT.*
21: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA2000.DAT.*
22: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA2001.DAT.*
23: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA2002.DAT.*
24: /SIDSI/gcgdata/geneseq/geneseq-emb1/AA2003.DAT.*
```

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

SUMMARIES

Result No.	Score	Query Match	Length	ID	Description
1	19	100.0	19	21	AA87840
2	19	100.0	19	23	AA871428
3	16	84.2	16	23	AA871430
4	15	78.9	15	23	AA871432
5	10	52.6	10	20	AA875078
6	10	52.6	11	21	AA879419
7	10	52.6	11	21	AA879547
8	10	52.6	11	22	AA875278
9	10	52.6	11	23	AA876085

10	10	52.6	11	24	ABP56078	Protein transducti
11	10	52.6	19	19	AAW41503	Heparin binding pe
12	10	52.6	19	21	AA87836	Heparin binding pe
13	10	52.6	19	23	AA871429	Peptide Bis-Arg He
14	9	47.4	11	20	AA875077	Transduction prote
15	9	47.4	11	21	AA829418	Synthetic transduc
16	9	47.4	11	21	AA829418	Amino acid sequenc
17	9	47.4	11	22	AA850377	Human immunodefici
18	9	47.4	11	23	ABG78988	Cell penetrating p
19	9	47.4	11	23	AAU76084	Peptide transport
20	9	47.4	11	23	AAU48625	Anti-inflammatory
21	9	47.4	11	23	AAU48626	Anti-inflammatory
22	9	47.4	11	24	ABP56077	Protein transducti
23	9	47.4	17	23	AAU48643	Anti-inflammatory
24	9	47.4	17	23	AAU48644	Anti-inflammatory
25	9	47.4	19	21	AAU78734	Heparin binding pe
26	9	47.4	22	23	AAU48636	Anti-inflammatory
27	9	47.4	22	23	AAU48637	Anti-inflammatory
28	9	47.4	92	20	AAU39179	M. tuberculosis an
29	9	47.4	92	20	AAU39036	M. tuberculosis re
30	9	47.4	105	23	ABU05688	M. tuberculosis an
31	9	47.4	160	20	AAU39186	M. tuberculosis an
32	9	47.4	160	20	AAU39043	M. tuberculosis re
33	9	47.4	190	24	ABP56096	PTD5-VP3 fusion pr
34	8	42.1	13	23	AAE23211	HIV-1 tat peptide
35	8	42.1	15	21	AAU08179	Peptide modulating
36	8	42.1	21	19	AAW41506	Heparin binding pe
37	8	42.1	21	21	AAU78739	Heparin binding pe
38	8	42.1	71	22	AAU46667	Propionibacterium
39	8	42.1	262	23	ABJ10474	Breast cancer - CA
40	8	42.1	262	23	ABJ10338	Novel human CASB74
41	8	42.1	262	23	AAE33614	Human CASB7439 pro
42	8	42.1	272	22	AAU49513	Propionibacterium
43	8	42.1	361	23	AAU10339	Novel human CASB74
44	8	42.1	450	24	ABJ37118	NOVX protein sequ
45	8	42.1	450	24	ABP57347	Human secreted pro

ALIGNMENTS

RESULT 1

AA87840

ID AA87840 standard; peptide; 19 AA.

XX AA87840;

AC AA87840;

XX 01-SEP-2000 (first entry)

DT Heparin binding peptide Bis-Arg helix #2.

XX Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

DE Heparin binding peptide; antagonist; cardiovascular; coagulant;

XX PS Example 1; Fig 1a; 39pp; English.

XX CC This invention describes novel heparin binding molecules (I). The

CC molecules (I) are useful as heparin antagonist drugs for cardiovascular

CC application and specifically neutralize heparin's conventional

CC anticoagulant properties. (I) are also useful for counteracting actions

CC of heparin locally e.g. in bleeding wounds, vascular anastomoses or

CC leaking prosthetic vascular grafts. (I) is also useful combined in a

CC pharmaceutical composition with insulin, as a substitute for protamine

CC for use in treating diabetics. The heparin binding molecules (I)

CC specifically neutralize heparin's conventional anticoagulant properties

CC without causing deleterious hemodynamic side-effects or exacerbation of

CC the proliferative vascular response to injury. (I) are short-duration,

CC intravenous drugs to be used in elective or emergency situations which

CC can safely and specifically neutralize heparin's proliferative response

CC to injury. This sequence represents a heparin-binding peptide described

CC in the method of the invention.

XX SQ Sequence 19 AA;

Query Match 100.0%; Score 19; DB 21; Length 19;

Best Local Similarity 100.0%; Pred. No. 3.6e-10;

Matches 19; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 ARAARRAARRAARRAARRA 19

DB 1 ARAARRAARRAARRAARRA 19

|||||

RESULT 2

AAB71428

ID AAB71428 standard; peptide; 19 AA.

AC AAB71428;

DT 27-NOV-2002 (first entry)

XX Peptide Bis-Arg Helix #2 fragment #1.

XX Sepsis; branched chain peptide; antibacterial; immunosuppressive;

KW endotoxin; helix peptide.

KX Synthetic.

OS

XX Key Location/Qualifiers

FT Modified-site 19

FT /note= "Ala is modified by unidentified R1 group"

XX

PN EP1232754-A2.

XX 21-AUG-2002.

PD 14-FEB-2002; 2002EP-0251027.

XX 14-FEB-2001; 2001US-268410P.

PR (COMM-) COMMONWEALTH BIOTECHNOLOGIES INC.

PA Harris RB, Wolz RL, Wolz G;

XX WPI; 2002-659478/71.

DR

XX Use of cationic helix peptides for treatment of sepsis and for the

PT detection and removal of endotoxins

PT Disclosure; Fig 1A; 18pp; English.

PS

XX This invention describes a novel use of antibacterial and

CC immunosuppressive peptides designated Arg Helix 2, Bis Arg Helix 2,

CC Tetra-Arg Helix 2 or Tris-Arg Helix 3 for the manufacture of a medicament

CC for the treatment of sepsis and the detection and removal of endotoxins.

CC The peptides of the invention are used in a method for detecting

CC endotoxin in a sample comprising contacting the sample with a labelled

CC helix peptide and then detecting the presence of any labelled molecule

CC bound to endotoxin. The peptides can also be used in a method for

CC removing endotoxin in a sample which comprises exposing the sample to a

CC helix peptide, bound to a solid support, then collecting the sample to a

CC endotoxin removal may be in vivo, or the peptides may be used to form an

CC affinity trap for endotoxins in e.g. dialysis-type treatments, or for

CC removal of endotoxins from plasma fractionation products. They are also

CC used as model frameworks for endotoxin binding from which new analogues

CC may be designed. This sequence represents the peptide Arg Helix #2 which

CC is used in the construction of Bis-Arg Helix #2, a branched chain peptide

CC described in the method of the invention.

XX SQ Sequence 19 AA;

Query Match 100.0%; Score 19; DB 23; Length 19;

Best Local Similarity 100.0%; Pred. No. 3.6e-10;

Matches 19; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 ARAARRAARRAARRAARRA 19

DB 1 ARAARRAARRAARRAARRA 19

|||||

RESULT 3

AAB71430

ID AAB71430 standard; peptide; 16 AA.

AC AAB71430;

DT 27-NOV-2002 (first entry)

XX Peptide Tris-Arg Helix #3 fragment.

XX Sepsis; branched chain peptide; antibacterial; immunosuppressive;

KW endotoxin; helix peptide.

KX Synthetic.

XX Key Location/Qualifiers

FT Modified-site 16

FT /note= "Ala is modified by unidentified R1 group"

XX EP1232754-A2.

XX 21-AUG-2002.

PD 14-FEB-2002; 2002EP-0251027.

XX 14-FEB-2001; 2001US-268410P.

PR (COMM-) COMMONWEALTH BIOTECHNOLOGIES INC.

PA Harris RB, Wolz RL, Wolz G;

XX WPI; 2002-659478/71.

DR

XX Use of cationic helix peptides for treatment of sepsis and for the

PT detection and removal of endotoxins

PT Disclosure; Fig 1B; 18pp; English.

PS

XX This invention describes a novel use of antibacterial and

CC immunosuppressive peptides designated Arg Helix 2, Bis Arg Helix 2,

CC Tetra-Arg Helix 2 or Tris-Arg Helix 3 for the manufacture of a medicament

CC for the treatment of sepsis and the detection and removal of endotoxins.

CC The peptides of the invention are used in a method for detecting

CC endotoxin in a sample comprising contacting the sample with a labelled

CC helix peptide and then detecting the presence of any labelled molecule

CC bound to endotoxin. The peptides can also be used in a method for

CC removing endotoxin in a sample which comprises exposing the sample to a

CC helix peptide, bound to a solid support, then collecting the sample. The

CC endotoxin removal may be in vivo, or the peptides may be used to form an

CC affinity trap for endotoxins in e.g. dialysis-type treatments, or for
 CC removal of endotoxins from plasma fractionation products. They are also
 CC used as model frameworks for endotoxin binding from which new analogues
 CC may be designed. This sequence represents the peptide Arg Helix #3 which
 CC is used in the construction of the peptide Arg Helix #3 which
 CC is used in the construction of Tris-Arg Helix #3, a branched chain
 CC peptide described in the method of the invention.

XX SQ Sequence 16 AA;
 Query Match 84.2%; Score 16; DB 23; Length 16;
 Best Local Similarity 100.0%; Pred. No. 1.2e-07;
 Matches 16; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
 Oy 4 ARRAARAARRARAEEA 19
 Db 1 ARRAARAARRARAEEA 16

RESULT 4
 AAB71432
 ID AAB71432 standard; peptide; 15 AA.

XX AC AAB71432;
 XX DT 27-NOV-2002 (first entry)
 XX DE Peptide Arg Helix #3 for construction of Tris-Arg helix #3.
 XX KW Sepsis; branched chain peptide; antibacterial; immunosuppressive;
 XX KW endotoxin; helix peptide.

XX OS Synthetic.
 XX FH Key Location/Qualifiers
 FT Modified-site 1 /note- "this residue has a side chain
 FT C(O)-NepsiloneH-(CH2)3-Tris-ArgHel#3, where
 FT the Tris-ArgHel#3 is represented in AAB71431"
 FT Modified-site 16 /note- "acylated residue"

XX PN EP1232754-A2.
 XX PD 21-AUG-2002.
 XX PF 14-FEB-2002; 2002EP-0251027.
 XX PR 14-FEB-2001; 2001US-268410P.

XX PA (COMM-) COMMONWEALTH BIOTECHNOLOGIES INC.

XX PI Harris RB, Wolz RL, Wolz G;

XX DR WPI; 2002-659478/71.

XX PT Use of cationic helix peptides for treatment of sepsis and for the
 XX detection and removal of endotoxins

XX PS Disclosure; Fig 2; 18pp; English.

XX CC This invention describes a novel use of antibacterial and
 CC immunosuppressive peptides designated Arg Helix 2, Bis Arg Helix 2,
 CC Tetra-Arg Helix 2 or Tris-Arg Helix 3 for the manufacture of a medicament
 CC for the treatment of sepsis and the detection and removal of endotoxins.
 CC The peptides of the invention are used in a method for detecting
 CC endotoxin in a sample comprising contacting the sample with a labelled
 CC helix peptide and then detecting the presence of any labelled molecule
 CC bound to endotoxin. The peptides can also be used in a method for
 CC removing endotoxin in a sample which comprises exposing the sample to a
 CC helix peptide, bound to a solid support, then collecting the sample. The
 CC endotoxin removal may be in vivo, or the peptides may be used to form an
 CC affinity trap for endotoxins in e.g. dialysis-type treatments, or for
 CC removal of endotoxins from plasma fractionation products. They are also

CC used as model frameworks for endotoxin binding from which new analogues
 CC may be designed. This sequence represents the peptide Arg Helix #3 which
 CC is used in the construction of the branched chain peptide Tris-Arg Helix
 CC #3 described in the method of the invention.

XX SQ Sequence 15 AA;

Query Match 78.9%; Score 15; DB 23; Length 15;
 Best Local Similarity 100.0%; Pred. No. 8.5e-07;
 Matches 15; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
 Oy 5 RRAARAARRARAEEA 19
 Db 1 RRAARAARRARAEEA 15

RESULT 5
 AAY25078
 ID AAY25078 standard; peptide; 11 AA.

XX AC AAY25078;

XX DT 24-AUG-1999 (first entry)
 XX DE Transduction protein peptide motif 3.

XX KW Anti-pathogen; fusion protein; protein transduction domain; PTD; A2T;
 XX KW cytotoxic domain; suppressor; infection; medicament; ddt; ddc; d4t; 3TC;
 XX KW FTC; DAPD; 1592089; CS92; acyclovir; ganciclovir; peniclovir; interferon;
 XX KW apoptosis; virus; HIV; cytomegalovirus; CMV; herpes simplex virus; HSV-1;
 XX KW hepatitis virus; Kaposi's sarcoma-associated herpes virus; KSHV;
 XX KW herpes virus; yellow fever virus; flavivirus; rhinovirus; plasmoidal;
 XX KW transduction efficiency; cytotoxin.

XX OS Unidentified.

XX PN WO9929721-A1.

XX PD 17-JUN-1999.

XX PF 10-DEC-1998; 98WO-US26358.

XX PR 20-APR-1998; 98US-0082402.

XX PR 10-DEC-1997; 97US-0069012.

XX PA (UNIW) UNIV WASHINGTON.

XX PI Dowdy SF;

XX DR WPI; 1999-394958/33.

XX PT New anti-pathogen systems, particularly for virus and plasmodium
 XX infections

XX PS Claim 69; Page 37; 123pp; English.

XX CC This invention describes a novel anti-pathogen system (APS) comprising a
 CC fusion protein constructed from a covalently linked protein transduction
 CC domain (PTD) and a cytotoxic domain. The APS can be used for suppressing
 CC a pathogen infection in a mammal. The method may further comprise
 CC administering a medicament e.g. AZT, ddt, ddc, d4t, 3TC, FTC, DAPD, The
 CC APS can also be administered to a mammal in the presence of a pathogen to
 CC induce apoptosis in a predetermined population of cells. The products can
 CC be used for treating mammals suffering from or susceptible to a viral
 CC infection or a disease associated with a virus, e.g. HIV, cytomegalovirus
 CC (CMV), herpes simplex virus, e.g. type 1 (HSV-1) hepatitis virus, type C
 CC (HCV), Kaposi's sarcoma-associated herpes virus (KSHV) or human herpes
 CC virus 8), yellow fever virus, flavivirus or rhinovirus, or suffering from
 CC or susceptible to plasmodial infection or a disease associated with a
 CC plasmodial infection, e.g. P. falciparum, P. vivax, P. ovale, or
 CC P. malariae. The APS exhibits high transduction efficiency and
 CC specifically kills or injures cells infected by one or more pathogens.

CC Formation of the cytotoxin is minimized or eliminated in uninfected cells
 CC and in infected cells that keep the pathogen inactive. The APS can be
 CC specifically tailored to kill or injure cells infected by one or more
 CC pathogen strains. This sequence represents a transduction protein motif
 CC described in the invention.

XX Sequence 11 AA;
 SQ Query Match 52.6%; Score 10; DB 20; Length 11;
 Best Local Similarity 100.0%; Pred. No. 0.014;
 Matches 10; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
 QY 1 ARAARRAARA 10
 Db 2 ARAARRAARA 11
 |||||

RESULT 6
 AAB29419
 ID AAB29419 standard; peptide; 11 AA.
 XX
 AC AAB29419;
 XX
 DT 09-FEB-2001 (first entry)
 XX
 DE Synthetic transduction peptide, SEQ ID NO:6.

XX Protein transduction domain; fusion molecule; therapeutic agent;
 KW drug targeting; drug discovery; cell transduction; bioavailability;
 KW vaccine; nervous system disorder; Alzheimer's disease;
 KW Parkinson's disease; Huntington's disease; pre-senile dementia; epilepsy;
 KW seizure; compulsive behaviour; meningitis; encephalitis; ischaemia;
 KW spongiform encephalopathy; dyslexia; age-related memory loss;
 KW Lou Gehring's disease; viral infection; HIV; bacterial infection.

XX Synthetic.
 OS
 XX WO200062067-A1.
 PN
 XX 19-OCT-2000.
 PD
 XX 28-FEB-2000; 2000WO-US05097.
 PF
 XX 28-FEB-1999; 99US-0122757.
 PR
 XX 29-AUG-1999; 99US-0151291.
 FR
 XX (UNIW) UNIV WASHINGTON.
 PA
 XX Dowdy SF;
 PI
 XX WPI; 2000-647439/62.
 DR
 XX

XX Fusion molecules comprising protein transduction domains and
 PT therapeutic agents, useful for treating e.g. Alzheimer's and
 PT Parkinson's diseases, dementia and epilepsy -
 PT
 XX Claim 36; Page 147; 191pp; English.

XX The invention relates to a novel fusion molecule comprising at least
 CC one protein transduction domain (PTD) and at least one linked molecule,
 CC where the linked molecule has therapeutic or prophylactic activity
 CC against a medical condition. The invention also relates to methods of
 CC drug discovery in which the test compound is linked to a suitable
 CC transducing protein and introduced to a cell; a method of killing
 CC resistant microorganisms using a suitable fusion molecule; a mammal
 CC comprising a covalently linked fusion molecule; and a mammal adapted for
 CC experimental use in which at least one transduction molecule has been
 CC transduced into essentially all the cells of the mammal. The fusion
 CC molecule is used to deliver a therapeutic agent to a mammal, especially
 CC a human. The linked molecule may be a vaccine, an anti-infective drug,
 CC a cardiovascular drug, an antitumour drug, an analgesic, an
 CC antiinflammatory, a diagnostic marker or a drug for the treatment or
 CC prevention of a central or peripheral nervous system disorder. The

CC central nervous system (CNS) disorder is especially Alzheimer's disease,
 CC Parkinson's disease, Huntington's disease, and also includes pre-senile
 CC dementia, epilepsy and seizures, compulsive behaviour, meningitis
 CC (including viral and bacterial meningitis), encephalitis, ischaemia,
 CC scrapie (or related spongiform encephalopathies), dyslexia, age-related
 CC memory loss or Lou Gehring's disease. Fusion molecules can also be
 CC used to kill virally infected cells, especially those infected with HIV.
 CC The vaccines are used to treat or prevent bacterial or viral infections.
 CC The methods are a highly effective means for transducing a molecule
 CC into an entire mammal or into specific cells, tissues, organs and
 CC systems within it. They also overcome bioavailability problems that
 CC are associated with many therapeutic agents (e.g., large molecular size,
 CC hydrophobicity, hydrophilicity, biological resistance), by providing
 CC efficient transduction of the target cell. The present sequence
 CC represents a specifically claimed protein transduction domain.

XX Sequence 11 AA;
 SQ Query Match 52.6%; Score 10; DB 21; Length 11;
 Best Local Similarity 100.0%; Pred. No. 0.014;
 Matches 10; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
 QY 1 ARAARRAARA 10
 Db 2 ARAARRAARA 11
 |||||

RESULT 7
 AAY93547
 ID AAY93547 standard; Peptide; 11 AA.
 XX
 AC AAY93547;
 XX
 DT 25-SEP-2000 (first entry)
 XX
 DE Amino acid sequence of a synthetic protein transduction domain.

XX Protein transduction system; protein transduction domain;
 KW cytotoxic domain; pathogen infection; retroviral infection;
 KW plasmoidal infection; cancer; prostate cancer.
 XX Synthetic.
 OS
 XX WO200034308-A2.
 PN
 XX 15-JUN-2000.
 PD
 XX 10-DEC-1999; 99WO-US9289.
 PF
 XX 10-DEC-1998; 98US-0111701.
 PR
 XX (UNIW) UNIV WASHINGTON.

PA Dowdy SF;
 PI
 XX WPI; 2000-431269/37.
 DR
 XX Protein transduction system for treating cancer and pathogenic
 PT infections has a fusion protein comprising a protein transduction
 PT domain covalently linked to a cytotoxic domain -
 XX Claim 69; Page 99; 127pp; English.

XX AAY93542-51 represent synthetic protein transduction domains, which
 CC are used in the protein transduction system of the invention. The
 CC specification describes a protein transduction system, which comprises
 CC a fusion protein. This fusion protein has a covalently linked protein
 CC transduction domain and cytotoxic domain. The system is useful for
 CC treating pathogen infection in mammals, infections such as those
 CC caused by CMV, HSV-1, HCV, KSHV, yellow fever virus, flavivirus or
 CC rhinovirus, retroviral infections such as HIV-1, HIV-2, HTLV-3 and/or
 CC LAV, plasmoidal infections associated with P.faciaparium, P.vivax,
 CC P.ovale, P.malariae. It is also useful for treating cancer, especially

SQ Sequence 11 AA;

Query Match 52.6%; Score 10; DB 23; Length 11;
Best Local Similarity 100.0%; Pred. No. 0.014;
Matches 10; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 ARAARRAARA 10
| | | | | | | | | |
DB 2 ARAARRAARA 11

RESULT 10

ASP56078
ID ABP56078 standard; Peptide; 11 AA.

AC ABP56078;

DT 27-FEB-2003 (first entry)

DE Protein transduction domain (PTD) peptide #4.

KW Cancer cell death; cancer; tumour; protein transduction domain; CAV;
KW chicken anaemia virus; cytostatic; proliferative cell disorder;
KW carcinogenesis; metastasis.

XX Unidentified.

OS WO200285305-A2.

PN 31-OCT-2002.

XX 24-APR-2002; 2002WO-US13092.

XX 24-APR-2001; 2001US-286099P.

XX (UNIW) UNIV WASHINGTON.

XX Dowdy SF, Ezhevsky SA, Wadla JS;

XX WPI; 2003-093056/08.

XX Novel fusion molecule useful for preventing or treating cancer,
PT comprises a protein transduction domain and a chicken anemia virus VP3
PT molecule -

XX Claim 24; Page 68; 104pp; English.

XX The present invention describes a fusion molecule (I) comprising at least
CC one protein transduction domain (PTD) and at least one chicken anemia
CC virus (CAV) VP3 molecule. (I) has cytostatic activity and can be used for
CC inducing cell death. (I) is useful for detecting cancerous or pre-
CC cancerous cells in a mammal or for killing or injuring cancerous or
CC pre-cancerous cells in a mammal. (I) is useful as a magnetic bullet to
CC selectively kill cancer cells in vitro and in vivo, for inducing cell
CC death, and for preventing or treating cancer and related proliferative
CC disorders. (I) is also useful for studying mechanisms of carcinogenesis
CC and metastases eukaryotic cells. (I) effectively transduces VP3 molecules
CC directly into the cells. (I) attacks cancer and pre-cancerous cells while
CC leaving normal cells relatively unharmed. Since more cells can be
CC targeted by (I) when compared with past attempts using different VP3
CC constructs, potential for patient relapse and side-effects are greatly
CC reduced. The present sequence represents a specifically claimed PTD
CC peptide which is given in the exemplification of the present invention.

XX Sequence 11 AA;

Query Match 52.6%; Score 10; DB 24; Length 11;
Best Local Similarity 100.0%; Pred. No. 0.014;
Matches 10; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 ARAARRAARA 10
| | | | | | | | | |
DB 2 ARAARRAARA 11

RESULT 11

AAW41503
ID AAW41503 standard; peptide; 19 AA.

XX AAW41503;

DT 05-JUN-1998 (first entry)

DE Heparin binding peptide.

KW Heparin binding peptide; anticoagulant antagonist; protamine;
KW insulin formulation; diabetes.

XX Synthetic.

XX WO9747312-A1.

XX 18-DEC-1997.

XX 03-JUN-1997; 97WO-US09037.

XX 11-JUN-1996; 96US-0660592.

XX (COMM-) COMMONWEALTH BIOTECHNOLOGIES INC.

XX Harris RB, Sobel M;

XX WPI; 1998-052023/05.

XX New peptide compounds - are useful as heparin binding molecules
XX which do not cause haemodynamic side effects

XX Claim 1; Page 43; 62pp; English.

XX The present heparin binding peptide can be used to antagonise or
CC neutralise the anticoagulant activity of heparin. It can also be
CC used to replace protamine in insulin formulations for
CC administration to diabetics.
CC The peptide can safely and specifically neutralise heparin's
CC anticoagulant properties, without causing deleterious haemodynamic
CC side-effects or exacerbating the proliferative vascular response to
CC injury.

XX Sequence 19 AA;

Query Match 52.6%; Score 10; DB 19; Length 19;
Best Local Similarity 100.0%; Pred. No. 0.021;
Matches 10; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 ARAARRAARA 10
| | | | | | | | | |
DB 10 ARAARRAARA 19

RESULT 12

AAV87836
ID AAV87836 standard; peptide; 19 AA.

XX AAV87836;

DT 01-SEP-2000 (first entry)

DE Heparin binding peptide Arg helix #2.

KW Heparin binding peptide; antagonist; cardiovascular; coagulant;
KW bleeding wound; vascular anastomoses; leaking prosthetic vascular graft;
KW protamine substitute; treatment.

XX Synthetic.

XX EP999219-A2.

```

XX PD 10-MAY-2000.
XX PF
XX PP
XX PR 01-OCT-1999; 99EP-0119514.
XX PR 06-OCT-1998; 98US-0166930.
XX PR (COMM-) COMMONWEALTH BIOTECHNOLOGIES INC.
XX PA
XX PI Harris RB, Sobel M;
XX PI WPI; 2000-306006/27.
XX DR
XX PT New heparin binding molecules, useful for reducing heparin content in a
XX PT mammal by reducing the anticoagulant effects of heparin -
XX PT
XX PS Example 1; Page 8; 39pp; English.
XX CC
XX CC This invention describes novel heparin binding molecules (I). The
XX CC molecules (I) are useful as heparin antagonist drugs for cardiovascular
XX CC application and specifically neutralize heparin's conventional
XX CC anticoagulant properties. (I) are also useful for counteracting actions
XX CC of heparin locally e.g. in bleeding wounds, vascular anastomoses or
XX CC leaking prosthetic vascular grafts. (I) is also useful combined in a
XX CC pharmaceutical composition with insulin, as a substitute for protamine
XX CC for use in treating diabetics. The heparin binding molecules (I)
XX CC specifically neutralize heparin's conventional anticoagulant properties
XX CC without causing deleterious hemodynamic side-effects or exacerbation of
XX CC the proliferative vascular response to injury. (I) are short-duration,
XX CC intravenous drugs to be used in elective or emergency situations which
XX CC can safely and specifically neutralize heparin's proliferative response
XX CC to injury. This sequence represents a heparin-binding peptide described
XX CC in the method of the invention.
XX SQ Sequence 19 AA;
XX
XX Query Match 52.6%; Score 10; DB 21; Length 19;
XX Best Local Similarity 100.0%; Pred. No. 0.021;
XX Matches 10; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
XX
QY 1 ARAARRAARA 10
DB 10 ARAARRAARA 19
|||||
|
RESULT 13
AAB71429
ID AAB71429 standard; peptide; 19 AA.
XX AC
XX AC AAB71429;
XX XX
XX DT 27-NOV-2002 (first entry)
XX DE
XX DE Peptide Bis-Arg Helix #2 fragment #2.
XX XX
XX KW Sepsis; branched chain peptide; antibacterial; immunosuppressive;
XX KW endotoxin; helix peptide.
XX OS
XX OS Synthetic.
XX XX
XX FH Key Location/Qualifiers
XX FT Modified-site 1
XX FT /note- "Ala is modified by unidentified R1 group"
XX XX
XX PN EP1232754-A2.
XX PD
XX PD 21-AUG-2002.
XX XX
XX PF 14-FEB-2002; 2002EP-0251027.
XX PR
XX PR 14-FEB-2001; 2001US-268410P.
XX PA (COMM-) COMMONWEALTH BIOTECHNOLOGIES INC.

```

```

XX Harris RB, Wolz RL, Wolz G;
XX WPI; 2002-659478/71.
XX Use of cationic helix peptides for treatment of sepsis and for the
XX detection and removal of endotoxins
XX PS Disclosure; Fig 1A; 18pp; English.
XX CC
XX CC This invention describes a novel use of antibacterial and
XX CC immunosuppressive peptides designated Arg Helix 2, Bis Arg Helix 2,
XX CC tetra-Arg Helix 2 or Tris-Arg Helix 3 for the manufacture of a medicament
XX CC for the treatment of sepsis and the detection and removal of endotoxins.
XX CC The peptides of the invention are used in a method for detecting
XX CC endotoxin in a sample comprising contacting the sample with a labelled
XX CC helix peptide and then detecting the presence of any labelled molecule
XX CC bound to endotoxin. The peptides can also be used in a method for
XX CC removing endotoxin in a sample which comprises exposing the sample to a
XX CC helix peptide, bound to a solid support, then collecting the sample. The
XX CC endotoxin removal may be in vivo, or the peptides may be used to form an
XX CC affinity trap for endotoxins in e.g. dialysis-type treatments, or for
XX CC removal of endotoxins from plasma fractionation products. They are also
XX CC used as model frameworks for endotoxin binding from which new analogues
XX CC may be designed. This sequence represents the peptide Arg Helix #2 which
XX CC is used in the construction of Bis-Arg Helix #2, a branched chain peptide
XX CC described in the method of the invention.
XX SQ Sequence 19 AA;
XX
XX Query Match 52.6%; Score 10; DB 23; Length 19;
XX Best Local Similarity 100.0%; Pred. No. 0.021;
XX Matches 10; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
XX
QY 1 ARAARRAARA 10
DB 10 ARAARRAARA 19
|||||
|
RESULT 14
AAY25077
ID AAY25077 standard; peptide; 11 AA.
XX AC
XX AC AAY25077;
XX XX
XX DT 24-AUG-1999 (first entry)
XX DE
XX DE Transduction protein peptide motif 2.
XX KW
XX KW Anti-pathogen; fusion protein; protein transduction domain; PTD; AZT;
XX KW cytotoxic domain; suppressor; infection; medicament; ddr; ddc; d47; 3TC;
XX KW FTC; DAPD; 1592089; CS92; acyclovir; ganciclovir; peniclovir; interferon;
XX KW apoptosis; virus; HIV; cytomegalovirus; CMV; herpes simplex virus; HSV-1;
XX KW hepatitis virus; Kaposi's sarcoma-associated herpes virus; KSHV;
XX KW herpes virus; yellow fever virus; flavivirus; rhinovirus; plasmodial;
XX KW transduction efficiency; cytotoxin.
XX OS
XX OS Unidentified.
XX PN WO9929721-A1.
XX PD
XX PD 17-JUN-1999.
XX XX
XX PF 10-DEC-1998; 98WO-US26358.
XX PR
XX PR 20-APR-1998; 98US-0082402.
XX PR 10-DEC-1997; 97US-0069012.
XX XX
XX PA (UNIW ) UNIV WASHINGTON.
XX XX
XX PI Dowdy SF;
XX WPI; 1999-394958/33.

```

```

XX PT New anti-pathogen systems, particularly for virus and plasmodium
XX PT infections
XX PS Claim 68; Page 37; 133pp; English.
XX CC This invention describes a novel anti-pathogen system (APS) comprising a
XX CC fusion protein constructed from a covalently linked protein transduction
XX CC domain (PTD) and a cytotoxic domain. The APS can be used for suppressing
XX CC a pathogen infection in a mammal. The method may further comprise
XX CC administering a medicament e.g. AZT, ddI, ddC, d4T, 3TC, FTC, DAPD,
XX CC 1592089, C892, acyclovir, ganciclovir, peniclovir or an interferon. The
XX CC APS can also be administered to a mammal in the presence of a pathogen to
XX CC induce apoptosis in a predetermined population of cells. The products can
XX CC be used for treating mammals suffering from or susceptible to a viral
XX CC infection or a disease associated with a virus, e.g. HIV, cytomegalovirus
XX CC (CMV), herpes simplex virus, e.g. type 1 (HSV-1) hepatitis virus, type C
XX CC virus 8), yellow fever virus, flavivirus or rhinovirus, or suffering from
XX CC or susceptible to plasmodial infection or a disease associated with a
XX CC plasmodial infection, e.g. P. falciparum, P. vivax, P. ovale, or
XX CC P. malariae. The APS exhibits high transduction efficiency and
XX CC specifically kills or injures cells infected by one or more pathogens.
XX CC Formation of the cytotoxic is minimized or eliminated in uninfected cells
XX CC and in infected cells that keep the pathogen inactive. The APS can be
XX CC specifically tailored to kill or injure cells infected by one or more
XX CC pathogen strains. This sequence represents a transduction protein motif
XX CC described in the invention.
XX CC Sequence 11 AA;
XX CC
XX CC Query Match 47.4%; Score 9; DB 20; Length 11;
XX CC Best Local Similarity 100.0%; Pred. No. 0.099;
XX CC Matches 9; Conservative 0; Mismatches 0; Indels 0; Gaps 0;
XX CC
XX CC QY 1 ARAARRAAR 9
XX CC | | | | | | | |
XX CC Db 2 ARAARRAAR 10
XX CC
XX CC RESULT 15
XX CC AAB29418
XX CC ID AAB29418 standard; peptide; 11 AA.
XX CC AC AAB29418;
XX CC
XX CC DT 09-FEB-2001 (first entry)
XX CC
XX CC DE Synthetic transduction peptide, SEQ ID NO:5.
XX CC
XX CC KW Protein transduction domain; fusion molecule; therapeutic agent;
XX CC KW drug targeting; drug discovery; cell transduction; bioavailability;
XX CC KW vaccine; nervous system disorder; Alzheimer's disease;
XX CC KW Parkinson's disease; Huntington's disease; pre-senile dementia; epilepsy;
XX CC KW seizure; compulsive behaviour; meningitis; encephalitis; ischaemia;
XX CC KW spongiform encephalopathy; dyslexia; age-related memory loss;
XX CC KW Lou Gehring's disease; viral infection; HIV; bacterial infection.
XX CC
XX CC OS Synthetic.
XX CC
XX CC PN WO200062067-A1.
XX CC
XX CC PD 19-OCT-2000.
XX CC
XX CC PF 28-FEB-2000; 2000WO-US05097.
XX CC
XX CC PR 28-FEB-1999; 99US-0122757.
XX CC PR 29-AUG-1999; 99US-0151291.
XX CC
XX CC PA (UNIV ) UNIV WASHINGTON.
XX CC
XX CC PI Dowdy SF;
XX CC

```

DR WPI; 2000-647439/62.

XX Fusion molecules comprising protein transduction domains and
 PT therapeutic agents, useful for treating e.g. Alzheimer's and
 PT Parkinson's diseases, dementia and epilepsy -

XX Claim 36; Page 147; 191pp; English.

XX The invention relates to a novel fusion molecule comprising at least
 CC one protein transduction domain (PTD) and at least one linked molecule,
 CC where the linked molecule has therapeutic or prophylactic activity
 CC against a medical condition. The invention also relates to methods of
 CC drug discovery in which the test compound is linked to a suitable
 CC transducing protein and introduced to a cell; a method of killing
 CC resistant microorganisms using a suitable fusion molecule; a mammal
 CC comprising a covalently linked fusion molecule; and a mammal adapted for
 CC experimental use in which at least one transduction molecule has been
 CC transduced into essentially all the cells of the mammal. The fusion
 CC molecule is used to deliver a therapeutic agent to a mammal, especially
 CC a human. The linked molecule may be a vaccine, an anti-infective drug,
 CC a cardiovascular drug, an antitumour drug, an analgesic, an
 CC antiinflammatory, a diagnostic marker or a drug for the treatment or
 CC prevention of a central or peripheral nervous system disorder. The
 CC central nervous system (CNS) disorder is especially Alzheimer's disease,
 CC Parkinson's disease, Huntington's disease, and also includes pre-senile
 CC dementia, epilepsy and seizures, compulsive behaviour, meningitis
 CC (including viral and bacterial meningitis), encephalitis, ischaemia,
 CC scrapie (or related spongiform encephalopathies), dyslexia, age-related
 CC memory loss or Lou Gehring's disease. Fusion molecules can also be
 CC used to kill virally infected cells, especially those infected with HIV.
 CC The vaccines are used to treat or prevent bacterial or viral infections.
 CC The methods are a highly effective means for transducing a molecule
 CC into an entire mammal or into specific cells, tissues, organs and
 CC systems within it. They also overcome bioavailability problems that
 CC are associated with many therapeutic agents (e.g., large molecular size,
 CC hydrophobicity, hydrophilicity, biological resistance), by providing
 CC efficient transduction of the target cell. The present sequence
 CC represents a specifically claimed protein transduction domain.

XX Sequence 11 AA;

Query Match 47.4%; Score 9; DB 21; Length 11;
 Best Local Similarity 100.0%; Pred. No. 0.099;
 Matches 9; Conservative 0; Mismatches 0; Indels 0; Gaps 0;

QY 1 ARAARRAAR 9
 | | | | | | | |
 Db 2 ARAARRAAR 10

Search completed: August 9, 2003, 16:29:05
 Job time : 54.7429 secs